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European Patent Office

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(11)

**EP 0 884 046 A1**

(12)

**EUROPEAN PATENT APPLICATION**

(43) Date of publication:

**16.12.1998 Bulletin 1998/51**

(51) Int. Cl.<sup>6</sup>: **A61K 7/48**

(21) Application number: **97201625.7**

(22) Date of filing: **30.05.1997**

(84) Designated Contracting States:

**AT BE CH DE DK ES FI FR GB GR IE IT LI LU MC  
NL PT SE**

Designated Extension States:

**AL LT LV RO SI**

(72) Inventors:

- **Eckart, Karl-Heinz Voss**  
**51375 Leverkusen (DE)**
- **Finkel, Peter**  
**51519 Odenthal (DE)**

(71) Applicant: **Sara Lee/DE N.V.**  
**3532 AA Utrecht (NL)**

(74) Representative:

**Smulders, Theodorus A.H.J., Ir. et al**  
**Vereenigde Octrooibureaux**  
**Nieuwe Parklaan 97**  
**2587 BN 's-Gravenhage (NL)**

(54) **Cosmetic composition with photoprotective properties**

(57) The invention is directed to a cosmetic composition with synergistically improved photoprotective properties, comprising

- at least one component selected from the group of Vitamin E, Vitamin C, derivatives of Vitamin E, derivatives of Vitamin C, and combinations of at least two of these components, and
- at least one natural polyamine or derivative.

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## Description

The invention is directed to cosmetic compositions with photoprotective properties, more in particular to compositions providing protection against UV-radiation from the sun (suncreams and the like)

UV exposure of the skin (UV-A to a greater extent than UV-B) leads to the generation of various free oxygen radicals in the skin. These free radicals attack in an entirely unselective manner precisely those skin constituents which are responsible for retention of elasticity and of moisture in the skin. Accordingly, they are starting point for several photochemical chain reactions, which finally may promote or lead to photo-induced skin damages like sunburn, skin ageing, immuno-suppression and possibly skin cancer.

Even if the skin is protected by UV-absorbing or -reflecting substances such as UV filters, pigments and micropigments, some UV radiation penetrates into the skin, which is quite often sufficient to start the above mentioned photochemical processes.

In order to provide a protection of the skin against the formation of these free radicals, skin-care compositions are provided on the skin which contain anti-oxidants that may penetrate the skin and act as radical scavengers.

Among the most effective radical scavengers are the Vitamin C and Vitamin E based derivatives. They may be used on the skin in various formulations, for example such as described in EP-A 579078. According to this publication the scavenger is used in combination with phytantriol, a compound enhancing the skin penetration.

However, there is a need for compositions having an improved effectivity as radical scavenger, either resulting in the use of smaller amounts of active components to obtain the same result, or in compositions having improved effectivity as photoprotective skin-care composition.

The invention is based on the surprising discovery, that naturally occurring polyamines, more in particular polyamines occurring in human or animal tissue (such as skin), in combination with Vitamin E and C, respectively derivatives thereof, show a synergistic improvement in effectivity.

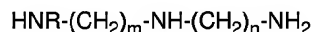
Accordingly the invention is directed to a cosmetic composition with photoprotective properties, comprising

- at least one component selected from the group of Vitamin E, Vitamin C, derivatives of Vitamin E, derivatives of Vitamin C, and combinations of at least two of these components, and
- at least one natural polyamine or derivative (such as a salt or an acid addition salt).

Surprisingly it has been found that a combination very small amounts of these polyamines (about 50 ppm)

with one or more of these Vitamins, provides a protection that is at least one order of magnitude larger than the protection of the Vitamin alone.

The naturally occurring polyamine is preferably a three- or tetra-amine, which contain at least one primary and at least one secondary amine group; more in particular it has the formula



wherein R denotes H or  $(\text{CH}_2)_k\text{NH}_2$ , wherein k, m and n are 2 or more, and the sum thereof does not exceed 10. Especially preferred compounds are spermine (m = 4, k = 3 and n = 3) and spermidine (R = H, m = 4 and n = 3).

The polyamine is present in an amount of 10 ppm, calculated on the weight of the composition, preferably in an amount of between 10 and 500 ppm.

The amounts of Vitamins to be used in the composition are preferably for Vitamin C or derivative thereof at least 0.005 wt.%, preferably between 0.005 and 1 wt.%, and for Vitamin E or derivative thereof at least 0.01 wt.%, preferably between 0.01 and 2.5 wt.%.

The composition according to the invention preferably contains both one component selected from Vitamin C and derivatives thereof, and one from Vitamin E and derivatives thereof. Such a composition provides an even better protection than systems based on polyamine with one Vitamin only.

Suitable derivatives of the Vitamins are salts and esters, such as the alkali and alkaline earth metal salts (sodium, potassium, calcium, magnesium and the like) and the phosphate, linoleate or acetate.

The composition further contains various conventional ingredients for photoprotective cosmetic compositions. Depending on the type of composition, body milk, suncream, sun lotion and the like various types of additives may be present.

As indicated above the use of phytantriol (2-(dihydroxyethyl)-2-hydroxy-6,10,14-trimethyl-pentadecane) or another vehicle for enhancing skin penetration may be contemplated. Suitable amounts will range from 0.05 to 20 wt.%, depending on the type of application.

The compositions generally contain at least one UV-filter in a preferred amount of between 0.05 to 15 wt.%. Suitable as UV filter are each and every UV-absorbing compounds mentioned in the EC positive list and which are published in the 14<sup>th</sup> Guideline 92/8/EEC of the European Commission, dated 18-2-92.

These filters are generally benzylidenecamphor compounds, p-aminobenzoic acid and derivatives thereof; benzophenone derivatives and benzotriazole derivatives.

The following compounds are preferably employed as UV filters:

N-propoxylated ethyl 4-aminobenzoate (mixture of isomers) ethoxylated ethyl 4-aminobenzoate glyceryl 4-aminobenzoate

2-ethylhexyl-4-dimethylaminobenzoate  
 2-ethylhexyl salicyclate  
 isopentyl 4-methoxycinnamate (mixture of isomers)  
 2-ethylhexyl 4-methoxycinnamate  
 2-hydroxy-4-methoxy-4'-methyl-benzophenone 5  
 [mexenone (INN)]  
 2-hydroxy-4-methoxybenzophenone-5-sulphonic  
 acid and sodium salt (sulisobenzone and sodium  
 salt),  
 $\alpha$ -(2-oxoborn-3-ylidene-toluene)-4-sulphonic acid 10  
 and salts thereof  
 3-(4'-methylbenzylidene)-d,1-camphor  
 3-benzylidenecamphor  
 4-isopropyl-dibenzoylmethane  
 4-isopropylbenzyl salicyclate 15  
 1-(4-tert-butylphenyl)-3-(4-methoxyphenyl)pro-  
 pane-1,3-dione  
 2,4,6-trianilin-(p-carbo-2'-ethylhexyl-1'-oxy)-1,3,5-  
 triazine.

The following compounds are particularly prefera-  
 bly used as UV filters:

2-ethoxyhexyl p-(dimethylamino)-benzoate;  
 2-ethylhexyl p-methoxycinnamate;  
 3-(4'-methylbenzylidene)-d,1-camphor;  
 2-hydroxy-5-methoxybenzophenone;  
 2-hydroxy-4-methoxybenzophenone-5-sulphonic  
 acid;  
 2-phenylbenzimidazole-5-sulphonic acid. 30

Along with the above-mentioned combination of  
 active compounds, the cosmetic product according to  
 the invention contains bases and auxiliaries which are  
 conventionally employed in cosmetics, in particular sta- 35  
 bilisers and antioxidants such as butylhydroxyanisole,  
 butylhydroxytoluene, EDTA salts such as magnesium  
 sulphate, in amounts from 0.02 to 5%, inter alia.

The bases and auxiliaries additionally include sol-  
 vents which are customary in cosmetics, such as water  
 to 80%, monoalcohols, lower polyalcohols having 1 to 6  
 carbon atoms or mixtures of these, furthermore fatty  
 material, such as mineral, animal, or vegetable oils such  
 as paraffin oil, or waxes such as microwax, fatty acids,  
 fatty alcohols, fatty acid esters such as cetylstearyl iso- 45  
 nonanoate and isopropyl palmitate, fatty alcohol ethers,  
 oxyethylated fatty alcohols, lanolin and derivatives, as  
 well as silicone oils in amounts from 0.5 to 50%, prefer-  
 ably 0.5 to 30%, particularly preferably in amounts from  
 5 to 30%.

If appropriate, the cosmetic skin-care product  
 according to the invention contains emulsifiers in  
 amounts from 0.1 to 20%, preferably in amounts from  
 0.2 to 10%, these emulsifiers being emulsifiers conven- 55  
 tionally employed in cosmetics, in particular non-ionic,  
 anionic, cationic or amphoteric compounds, for example  
 sterols, polyol fatty acid esters and polyol fatty alcohol  
 ethers, alkali metal salts and triethanolamine salts of

fatty acid, sodium cetylstearyl sulphate, tetracylammo-  
 nium halides and phospholipids. Examples of these are  
 glycerol sorbitan fatty acid esters, polyoxyethylene fatty  
 acid esters and alkyltetraglycol ether o-phosphoric acid  
 esters.

0.02 to 5%, preferably 0.1 to 2%, of thickeners and  
 gelling agents can furthermore be employed in the  
 product according to the invention. These include poly-  
 acrylic acid derivatives, cellulose derivatives, ben-  
 tonites, xanthan derivatives, alginates, guar gum and  
 locust bean gum Polyacrylamide and zinc stearate are  
 examples.

The preparation according to the invention can con-  
 tain other substances which are customary in cosmet-  
 ics. These include humectants (0.5 to 15%), colorants,  
 buffer substances, preservatives and perfume oils in  
 amounts from 0.01 and 5.0%.

The following may be mentioned by way of example  
 as humectants: lower polyalcohols such as glycerol,  
 propylene glycol, butylene glycol, sorbitol, moreover 2-  
 pyrrolidone-5-carboxylic acid and the sodium salt  
 thereof, lactic acid and the salts thereof, urea, proteins  
 and protein derivatives such as collagent, and further-  
 more hyaluronic acid, inter alia.

Colorants to be added to the cosmetic preparations  
 according to the invention which may be mentioned by  
 way of example are:

Colour C.I. 16255, colour C.I. 61570, colour C.I.  
 42051, colour C.I. 15985, colour C.I. 77492. The follow-  
 ing are preferably suitable as preservatives:

2,4-hexadienoic acid (sorbic acid and salts thereof),  
 4-hydroxybenzoic acid, salts and esters thereof,  
 3-acetyl-6-methyl-2,4(3H)-pyrandione (dehydrace- 35  
 tic acid) and salts thereof,  
 1,1-methylene-bis-(3-(1-hydroxymethyl-2,4-diox-  
 imidazolidin-5-yl)-urea),  
 imidazolindinyurea,  
 2-phenoxy-ethanol,  
 benzyl alcohol. 40

The cosmetic skin-care product according to the  
 invention is preferably in the form of an emulsion (cream  
 or milk), such as oil-in-water or water-in-oil emulsions. It  
 is generally prepared by mixing and stirring of the com-  
 ponents, if appropriate followed by homogenisation, and  
 if appropriate and preferably in an evacuated apparatus.

All percentages in the present text relate to percent-  
 ages by weight, unless stated otherwise.

The invention is illustrated hereinafter in greater  
 detail with the aid of the examples, without this being  
 intended to have a restrictive character.

#### EXAMPLE 1

By the use of the in-vivo  $\beta$ -carotin model:

- dermal application of the test product

- waiting period ensuring penetration of the formulation into skin
- treatment with  $\beta$ -carotin
- irradiation of the skin with UV light containing sufficient UV-A (test area and untreated control)
- compare photochemical degradation of  $\beta$ -carotin by UV-induced radicals of treated and untreated skin

To achieve a sufficient protection level about 0.6 % Vitamin E or 0.1 % of Vitamin C had to be used. In combination with 50 ppm spermine the level of Vitamin C could be 0.05 % and of Vitamin E 0.1 %, for obtaining the same level of activity.

A combination of 50 to 100 ppm spermine, 0.05 % Vitamin E and 0.01 % Vitamin C also provided sufficient protection.

## EXAMPLE 2

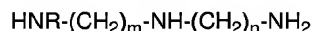
A sun protection balm was prepared from:

Glycerol (waterfree)	3.0
D-panthenol	1.0
Xanthan gum	0.1
carbomer	0.5
45 % NaOH	0.45
Vitamin C phosphate	
Sodium salt	0.1
Spermine	0.025
96% ethanol	8.0
Isopropylpalmitate	6.0
phenyl trimethicone	3.0
Tocopherolacetate	1.0
Phytantriol	0.1
Perfume	0.5
Water	76.225

## Claims

1. Cosmetic composition with photoprotective properties, comprising
  - at least one component selected from the group of Vitamin E, Vitamin C, derivatives of Vitamin E, derivatives of Vitamin C, and combinations of at least two of these components, and
  - at least one natural polyamine or derivative.

2. Composition according to claim 1, wherein the composition additionally contains UV-absorbing and/or reflecting compounds.
3. Composition according to claim 1 or 2, wherein the composition additionally contains one or conventional ingredients for cosmetic photoprotective compositions.
4. Composition according to claim 1-3, wherein the composition contains a polyamine naturally occurring in human or animal tissue, as the natural polyamine.
5. Composition according to claim 4, wherein the natural polyamine is a three- or tetra-amine, which contain at least one primary and at least one secondary amine group.
6. Composition according to claim 5, wherein the polyamine has the formula



wherein R denotes H or  $(\text{CH}_2)_k\text{NH}_2$ , wherein k, m and n are 2 or more, and the sum thereof does not exceed 10.

7. Composition according to claim 1-6, wherein the natural polyamine is present in an amount of at least 10 ppm, calculated on the weight of the composition, preferably in an amount of between 10 and 500 ppm.
8. Composition according to claim 1-7, wherein a combination of at least one of Vitamin E or derivative of Vitamin E, and Vitamin C or, derivative of Vitamin C, is present.
9. Composition according to claim 1-8, wherein the amount of Vitamin C or derivative thereof is at least 0.005 wt.%, preferably between 0.005 and 1 wt.%, and the amount of Vitamin E or derivative thereof is at least 0.01 wt.%, preferably between 0.01 and 2.5 wt.%.
  - 10. Composition according to claim 1-9, wherein the derivatives of Vitamin E and Vitamin C are selected from salts and esters thereof.
  - 11. Composition according to claim 1-10, in the form of a sun lotion, body milk or suncream.
  - 12. Use of a combination of a natural polyamine and at least one of Vitamin E, Vitamin C, derivatives of Vitamin E and derivatives of Vitamin C, for synergistically improving the UV-protection level in cosmetic compositions.



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# EUROPEAN SEARCH REPORT

Application Number  
EP 97 20 1625

DOCUMENTS CONSIDERED TO BE RELEVANT			
Category	Citation of document with indication, where appropriate, of relevant passages	Relevant to claim	CLASSIFICATION OF THE APPLICATION (Int.Cl.6)
X	DATABASE WPI Week 9550 Derwent Publications Ltd., London, GB; AN 95-390534 XP002046837 "Antioxidant for pharmaceuticals, feed and cosmetics-contains spermine, spermidine and/or putrescine and oil and fat" * abstract * & JP 07 268 323 A (NISSHIN OIL MILLS LTD) 17 October 1995 ---	1,3-12	A61K7/48
X	DATABASE WPI Week 9551 Derwent Publications Ltd., London, GB; AN 95-400854 XP002046838 "cosmetic materials with high oxidation stability-contain fat and oil composition containing spermine, spermidine and/or putrescine" * abstract * & JP 07 277 917 A (NISSHIN OIL MILLS LTD) 24 October 1995 ---	1,3-8, 10-12	TECHNICAL FIELDS SEARCHED (Int.Cl.6) A61K
A	M-C. MARTINI: "Actifs et Additifs en Cosmétologie" 1993, LAVOISIER TEC & DOC., PARIS (FR) XP002046836 * page 227-228 * ---	1-12	
A	WO 96 23490 A (COSMEDERM TECHNOLOGIES ;HAHN GARY SCOTT (US); THUESON DAVID OREL) () * claims 1,2,18,37,109 * -----	1-12	
The present search report has been drawn up for all claims			
Place of search THE HAGUE		Date of completion of the search 13 November 1997	Examiner Sierra Gonzalez, M
CATEGORY OF CITED DOCUMENTS X : particularly relevant if taken alone Y : particularly relevant if combined with another document of the same category A : technological background O : non-written disclosure P : intermediate document		T : theory or principle underlying the invention E : earlier patent document, but published on, or after the filing date D : document cited in the application L : document cited for other reasons & : member of the same patent family, corresponding document	

EPO FORM 1503 03/82 (P4/C01)